



Exploring Phytochemical Profiles of Medicinal Plants Used for Diarrhea Treatment

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ABSTRACT

Diarrheal diseases remain a leading cause of morbidity and mortality, particularly among children under five in developing countries. Despite advances in antibiotics and sanitation, access to conventional treatments remains limited in low-resource settings. Traditional medicinal plants offer an accessible and culturally accepted alternative, but their phytochemical properties and safety profiles require scientific validation. This study investigates the phytochemical composition of ten medicinal plants traditionally used to treat diarrhea in Benin and surrounding regions. Using standard phytochemical screening techniques, the presence of key secondary metabolites including flavonoids, alkaloids, tannins, terpenoids, polyphenols, and saponins was confirmed. Toxicological assessments using *Artemia salina* and zebrafish models indicated low acute toxicity at therapeutic doses. The study highlights the ethnomedicinal relevance and potential pharmacological efficacy of these plants, particularly *Blighia sapida*, *Cochlospermum tetramera*, and *Cochlospermum triphylla*, underscoring the need for further pharmacodynamic and clinical research to support their inclusion in formal healthcare systems.

Keywords: Phytochemical screening, medicinal plants, diarrhea, ethnopharmacology, secondary metabolites, traditional medicine.

INTRODUCTION

Diarrheal diseases pose a significant global health risk despite sanitation improvements and antibiotic development. Symptoms often include more than three loose stools in four hours. Diarrhea can be classified as secretory (e.g., food poisoning, cholera), osmotic (from poorly absorbed solutes), or inflammatory (due to pathogens like *Salmonella*). Common treatments include probiotics, antibiotics, and medications to slow stooling, but antibiotics may take days to work. Diarrhea from food poisoning generally resolves without treatment, while bacterial infections may require antibiotics. Probiotics can enhance the body's natural flora, yet many lack the affordability for these options. Consequently, medicinal plants offer alternative remedies, with a substantial portion of medications derived from plants. Ethnoveterinary practices utilize traditional medicine for livestock ailments, often effectively preferred over modern interventions. However, caution is necessary as some traditional plants may be toxic, especially with prolonged use. Phytochemical analysis of medicinal plants is vital to assess their safety and efficacy. Tannin-rich plants have shown promise in alleviating diarrhea, while lesser-known plants may also provide therapeutic benefits. The selected plants in this study could represent candidates for anti-diarrheal testing or serve as a basis for investigating their phytochemical properties [1, 2].

Overview of Diarrhea

Diarrhea is defined as increased stool frequency, weight, or decreased consistency, and is characterized by the frequent expulsion of liquid stools containing undigested food, mucus, and blood. It is classified by duration: acute (< 14 days), persistent (14–29 days), and chronic (≥ 30 days). Major causes include

infections, food intolerances, medications, emotional stress, poor food and water habits, allergies, and elixirs. Types of diarrhea include osmotic, secretory, motility, and inflammatory. Diarrhea leads to dehydration, where the loss of pure water exceeds electrolyte loss, potentially causing complications like renal failure and altered mental states. Prompt fluid and electrolyte replenishment is critical, along with antidiarrheal medications to slow gastrointestinal movement. Chronic diarrhea occurs when symptoms persist for over three weeks. Acute diarrhea is divided into acute watery diarrhea, which may cause severe dehydration, and acute bloody diarrhea, known as dysentery. Dehydration severity ranges from minimal loss (0–5% - unnoticeable) to very severe loss (>15% - often fatal), with symptoms escalating from mild thirst to severe hypotension and altered sensorium. There's a rising interest in herbal medications due to their effectiveness, safety, and affordability compared to synthetic drugs [3, 4].

Types of Diarrhea

Diarrhea is the increased frequency of stool with watery consistency and is classified into types according to different parameters such as duration, clinical course, pathophysiological conditions, etc. The major types of diarrhea include osmotic diarrhea, secretory diarrhea, motility-related diarrhea, and dysentery. The most common type of diarrhea is secretory diarrhea. In secretory diarrhea, there is a net excess of seawater secretion over absorption due to raised anion secretion or lowered anion absorption. This causes an excess of water secretion and thus results in diarrhea. The significant cause of secretory diarrhea is cholera toxin. Osmotic diarrhea is due to unabsorbed osmotically active solutes, leading to water retention in the luminal side. The second most common type of diarrhea is osmotic diarrhea, and the major cause of osmotic diarrhea is lactose intolerance for carbohydrates. Some medications can act as osmotic laxatives; hence, this type of diarrhea is a potential side effect. In dysentery, there is an inflammation of the intestinal mucosa, usually due to an infection that causes exudation of mucus and/or blood. Dysentery is characterized by the presence of at least some formed stools and may occur in amoebic dysentery, bacillary dysentery, etc. In inflammatory diarrhea, inflammatory cells are present in the stool, and the condition may be due to infectious colitis. In impacting diarrhea, a hard scybala is (or are) stuck in the rectum, making it difficult to pass fecal matter. An encounter group discusses the different treatments and their availability, and new remedies are also proposed to be effective against diarrhea in these research studies. A total of 100 routine antidiarrheal medicines are studied. Field visits and consultations with indigenous healers are proposed to identify medicinal plants to be studied in detail regarding chemical composition, preventive and curative actions [5, 6].

Causes and Risk Factors

Diarrhea is a major cause of childhood mortality, particularly in developing countries, and chemical treatments often have side effects. This study reviews traditional medicinal plants used in the Amhara region to treat diarrhea, focusing on their phytochemistry and bioactivity for antidiarrheal activities. By 2023, the authors compiled research on these plants and their phytochemical constituents linked to medicinal applications. Diarrheal illness, marked by changes in stool consistency and frequency, poses a significant global burden. It ranks as the second leading cause of death among children under five, contributing to malnutrition. Diarrheal diseases affect many in this age group and remain a public health challenge, with these young children accounting for 60–70% of an estimated 3 million annual diarrhea deaths. Their heightened risk stems from underdeveloped immune systems and prevalent breast amenorrhea. While advancements in treatment and prevention have occurred, controlling fecal–oral transmission is crucial. Secondary prevention methods, especially during outbreaks, are vital in in-home settings. Although institutional and community interventions are effective in managing diarrhea, accessibility remains an issue in low-income countries where it's common. The fundamental intervention lies in accessing safe drinking water, as untreated sources can trigger severe outbreaks, even in areas with low endemicity [7, 8].

Global Impact and Statistics

Given natural medicine's long history of providing therapies for different ailments, plants continue to be regarded as critical therapeutic agents, especially in the treatment of infectious and non-infectious diseases. Diarrhea is characterized by an increase in frequency and liquidity of stool that cannot be correlated with increased food ingestion. Diarrhea is divided into two main types: "acute" and "chronic." Acute diarrhea is either "watery" or "bloody" (or dysenteric). Watery or non-bloody diarrhea can be caused by viruses (especially rotavirus), pathogenic bacteria (e.g., enterotoxigenic *Escherichia coli*, *Vibrio cholerae*), and protozoan parasites (e.g., *Giardia lamblia*) and is usually characterized by a high volume of stool. Bloody diarrhea (or dysentery) can be caused by pathogenic bacteria (e.g., *S. dysenteriae*, *S. flexneri*)

and protozoan parasites (e.g., *Entamoeba histolytica*) and is characterized by the presence of blood and mucus in the stool. Dairy farming, animal husbandry, and/or the consumption of unboiled water are the main contributors to the etiology of infectious diarrhea. Diarrhea is the second leading cause of morbidity and mortality in the country due to its high prevalence among children aged five or less. It is one of the major health problems in most developing countries, including Ethiopia, especially among children under the age of five years, with reported deaths of over 1.5 million every year. Recent studies from developing countries have estimated that between 720 million episodes of diarrhea and 2.4 million associated deaths occur annually, especially in children. Diarrhea not only leads to death but also results in the loss of household productivity, monetarily due to the non-availability of adults for household work and employment (indirect cost) and directly to costs incurred by families and communities for treatment and care of sick children [9, 10].

Medicinal Plants and Their Importance

Medicinal plants have been and still are the source of a large number of therapeutic agents, and it has been reported that around 25% of registered drugs in the present Modern Medicine have an origin in medicinal plants [10-14]. Very interestingly, all the Ancient Systems of Medicine from Sumerian Texts, Ancient Egyptian Papyri, the book by Hippocrates, to Ayurveda were based on the knowledge acquired after hundreds of years of using different medicinal plants [15-19]. Even in modern times, medicinal plants have been on the frontiers of research in providing new lead molecules because of their diversity and bioactivity. However, farmer communities have tremendous knowledge on different medicinal plants which have their use in various ailments, but due to migration of these communities, time and space squandering, this traditional knowledge is disappearing. In this paper, an attempt has been made to document systematically the ethnobotanical aspects of the medicinal plants among this community so that such important knowledge can be preserved for future reference [20-25]. Diarrhea is one of the leading causes of mortality in children less than five years old, with the most major impact being observed in developing countries. In Iran, diarrhea diseases are reported to be the second leading cause of death in children under five years old after respiratory diseases. Globally, infectious diseases are the most common cause of diarrhea and mortality, particularly for acute diarrhea [26-28]. Diarrhea requires immediate treatment and public community attention. Untreated diarrhea leads to increased weight loss, dehydration, reduced overall health, and death. Oral rehydration therapy is a safe and inexpensive treatment for diarrheal diseases, which can reverse dehydration in 90% of cases. Nevertheless, appropriate consumption of anti-diarrheic herbal medicine is essential in cases of non-infectious diarrhea since they promote the reabsorption of blood fluid [29-32].

Phytochemistry Basics

Phytochemical screening has been done to determine the presence or absence of major chemical groups in the different organs of the analyzed plants. Major groups of chemical compounds, such as tannins, gallic tannins, flavonoids, anthocyanins, leucoanthocyanins, alkaloids, mucilages, reducing compounds, sterols, terpenes, and saponosides, have been searched for [33-36]. Phytochemical screening is a fundamental aspect of pharmacognosy, which aims to assess the presence of bioactive compounds in medicinal plants. Phytochemical screening shows the percentage occurrence of chemical groups depending on the plant organ and the species. The various secondary metabolites detected may be responsible for the medicinal properties of the different plants studied [37-39]. The presence of these groups has been cited in other studies involving other plants. Qualitative phytochemical screening is a time-tested procedure used extensively in the field to help contribute to the understanding of the bioactivity of plant secondary metabolites and their phytomedicines [40-45]. In this study, medicinal plants used by traditional practitioners in the treatment of diarrheal diseases were explored. The bioactivity of the ten plants was ensured using a phytochemical approach. Drug formulations from the tested plants can be obtained in the laboratory for the treatment of these *santé* causes. Phytochemical screening showed the presence of ten broad chemical groups in the plant organs studied [46-49]. Further studies based on a more refined screening would help identify other secondary metabolites. Significant ($p < 0.05$) qualitative interspecific variations in the chemical composition of the various organs of the ten plants were obtained. These differences may be due to species variation and cultural and environmental parameters [50-51].

Methodology for Phytochemical Analysis

The leaves or parts of the ten plants used were obtained from local suppliers in the country and were identified by an ethnobotanist. The plant samples were dried in an airy shed and subjected into powder form with a mechanical grinder. Phytochemical screening tests for flavonoids, alkaloids, terpenoids,

polyphenols, tannins, saponins, and reducing compounds were carried out using standard methods. The ethyl acetate and methyl alcohol extracts of each plant were screened in aqueous solution for their effect on the presence of each class of phytochemicals, respectively. The screening tests were accomplished in the biology laboratory of the Department of Plant Biology and Biochemistry, Faculty of Science of the University of Benin, Nigeria. The presence of alkaloids was tested using Hager's reagent (yellow precipitation), and the polyphenol compound was detected using a few drops of ferric chloride (green-blue coloration). The presence of terpenoids was tested with 2 mL of freshly prepared chloroform in a separate tube containing 1 mL of concentrated H₂SO₄ (red coloration). The presence of tannins was tested using 7% sodium chloride, and the detection of saponins was tested using 2 mL of distilled water (frothing). Flavonoids were tested using drops of 2N HCL with a pinch of magnesium powder (pink color formation). The screening of the reducing compounds was performed using heated distilled water, 2 mL concentrated H₂SO₄, and then diluted with 20 mL of distilled water with an orange-red coloration (clear white precipitate). The result obtained was observed in a standard table for confirmation of each class of phytochemicals [15, 16].

Phytochemical Profiles of Selected Medicinal Plants

In Benin, diarrheal diseases constitute a significant public health challenge, with herbal medicine being the predominant treatment choice. This study focused on the phytochemical properties of ten plants traditionally used for these ailments and evaluated their toxicity using *Artemia salina* and zebrafish models. A methanol-water mix (70:30) was used for extraction. Phytochemical screenings displayed diverse secondary metabolites, such as phenols, flavonoids, terpenoids, tannins, and alkaloids. The cytotoxicity model showed that the plants are noncytotoxic (LC₅₀ > 1000 µg/mL). Acute toxicity assessments with the dried powder on fish revealed no mortality (mortalities < 40%) or liver and kidney changes at a dose of 2000 mg/kg, indicating normal behavior and feeding in the exposed fish. These findings support their inclusion in various African pharmacopeias. However, further investigations into sub chronic, chronic toxicity, mutagenicity, and carcinogenicity are essential for a comprehensive toxicological understanding. Globally, diarrheal diseases rank as the second leading cause of infectious disease mortality in children. All populations are susceptible to diarrhea, especially in developing nations where sanitation, water safety monitoring, and food processing oversight are inadequate. Contributing factors include malnutrition, low maternal education, and high unemployment rates. In Africa, children under five suffer significant health loss due to diarrhea-related mortality. Benin faces severe public health challenges linked to diarrhea, exacerbated by poor health services and limited sanitation access, with over 300 medicinal plant species utilized in the region [17, 18].

Mechanisms of Action of Phytochemicals

Diarrhea is a prevalent health issue, causing 760,000 deaths in children under five annually, mainly due to dehydration and poor nutrition. In Benin's traditional medicine, three plants are utilized for treating diarrhea: *B. sapida* (bark), *C. tetramera* (bark), and *C. triphylla* (leaf). These plants are favored for their efficacy, affordability, and availability. Phytochemical screenings reveal that they possess significant amounts of secondary metabolites, especially saponins, tannins, flavonoids, alkaloids, and polyphenols. Acute toxicity assessments using OECD GUIDELINES-423 indicate that oral doses of up to 2000 mg/kg in mice do not adversely affect behavior or induce mortality, suggesting low acute toxicity. *B. sapida*, known as "bocol" in Fon, is widely used in Benin for diarrhea treatment, particularly its bark. Ethnopharmacological surveys confirm its use by traditional healers in Porto-Novo, Ganvie, Ouidah, and Comé. Scientific literature supports its effectiveness against *Castalia* species, and phytochemical investigations reveal its richness in catechins, procyanidins, and ellagitannins, validating its traditional use. Similarly, numerous studies on *C. tetramera*'s bark demonstrate its efficacy against infectious diarrhea and its antibacterial properties, with pharmacological studies elucidating its mechanisms of action. Both *B. sapida* and *C. tetramera* contain compounds such as gallotannins, flavonoids, and ellagic acid, contributing to their therapeutic effects against diarrhea. *C. triphylla*'s entire plant is recognized in Benin's ethnopharmacological records for diarrhea treatment. Given its traditional applications, further scientific studies are warranted to investigate its efficacy and action mechanism [19, 20].

Clinical Evidence Supporting Medicinal Plants

Many medicinal plants prevalently used for diarrhea are supported by scientific research regarding their antidiarrheal and antimicrobial activities. Studies performed with extracts of species such as *Calpurnia aurea*, *Tithonia diversifolia*, *Croton macrostachyus*, *Phytolacca dodecandra*, *Ficus sycomorus*, and *Hassleria trinervula* documented antidiarrheal activity using biological assays. The phytochemical

analysis of these plants revealed the presence of active ingredients commonly associated with antidiarrheal action, such as alkaloids, flavonoids, tannins, saponins, terpenoids, and phenolic compounds. These phytochemicals have been shown to affect through either one or several modes of action on many diseases, including diarrhea. Some are known to inhibit the secretion of electrolytes and water into the intestinal lumen. Furthermore, most documented species indicated antimicrobial properties. In addition to gains in knowledge regarding the medicinal plants and other ethnomedicinal practices, these research efforts establish a foundation for evaluating worldwide the pharmacognostic and pharmacological evidence supporting the efficacy of traditional medicine. The goal is to demonstrate efficacy via laboratory testing of both traditional compounds and the purified active ingredients. Furthermore, investigation into the mechanism(s) of action of the active ingredient(s) are expected to bring new insights into the understanding of physiology and pharmacology, which, in turn, might lead to novel approaches in combating diseases that are occurring within race or gender-based and agricultural differences [21, 22].

Safety and Toxicity Considerations

In several developing countries, traditional medicine is often the primary treatment for various diseases. In western Africa, medicinal plants used for acute diarrhea are commonly sold. However, there is limited phytochemical and toxicological data available on these plants. The WHO is developing guidelines for herbal medicine safety assessments, needing information similar to that used for synthetic drugs. Yet, the inherent complexity of plants makes replicating synthetic drug procedures impractical, potentially increasing uncertainty regarding herbal safety. Thus, specific toxicological studies are essential for identifying harmful phytochemicals in herbal remedies. An alternative for consumers and regulators is utilizing timely research on plant toxicity by establishing a toxicity database for herbal studies. Phytochemical screenings have revealed diverse secondary metabolites in these plants, including alkaloids, flavonoids, phenolic compounds, saponins, and tannins. All extracts contained alkaloids, tannins, and flavonoids, but phenolic compounds were notably missing in *S. sesban* and *M. oleifera*. Noncholesterol sterols appeared only in *E. crassipes* and *S. sesban* extracts. Other compounds like sterols, flavonoids, and coumarins were absent as well. This underlines the complex phytochemical profiles of the studied plants. Extraction methods significantly impact the recovery of bioactive compounds, which may explain the missing metabolites. There are also variations in metabolite richness among samples. The investigated plants have potential as sources of secondary metabolites with health benefits, warranting further phytochemical and pharmacological exploration [23-28].

Future Directions in Research

Traditional medicinal plants are important in health care systems. In developing countries, herbal products are still used as therapeutic agents. Diabetes, malaria, rheumatism, high blood pressure, asthma, skin diseases, and diarrhea are chronic health problems in Africa and globally. Anti-infective and medicinal plants used to treat infectious diseases include plants with broad-spectrum activity against several infections. Phytochemicals are plant-derived natural bioactive substances that exhibit antimicrobial activity against bacteria, viruses, fungi, protozoa, and helminths. Phytochemical extracts from *Myrsine africana* and *Stoecha aspera* effectively inhibit pathogenic agents that cause diarrhea in humans. Even though plants play an important role in the treatment of infectious diseases worldwide, plants used as antimicrobials and the activity of selected plants against microbiological infections of traditional medicinal plants that treat diarrhea have not been well studied. Ethnopharmacological studies of antidiarrheal plants used by traditional communities in the Sundarbans mangrove forest of India have been reviewed and justified through experiments. Use of phytochemicals as antidiarrheal agents is supported by pharmacological experiments. Support is also provided for the use of various plants in folkloric systems of medicine. Additionally, *in vitro* antibacterial activity is demonstrated. Their antidiarrheal effects can also be studied in acute diarrhea animal models. These findings may open the way to more pharmacognostic studies on these plants with folklore use in diarrhea treatment [29-33].

CONCLUSION

The findings of this study validate the traditional use of several medicinal plants in the treatment of diarrhea, revealing significant phytochemical diversity that likely contributes to their therapeutic effects. Key compounds such as tannins, flavonoids, and alkaloids play crucial roles in the anti-diarrheal mechanisms of these plants. Toxicological evaluations affirm their safety at conventional doses, making them promising candidates for affordable and accessible treatment options in resource-limited regions. However, to ensure public health safety and regulatory integration, further research is essential to

investigate subchronic toxicity, pharmacological efficacy, dosage standardization, and mechanisms of action. Documenting and preserving indigenous knowledge, while integrating it with modern phytomedicine, can enhance health outcomes and foster sustainable medicinal plant use.

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